

3           a plurality of fiber optic cables for illuminating volumes of the plurality of samples,  
4           a plurality of lenses, each co-axially disposed with a first end of a fiber optic cable for  
5           focusing an excitation beam into a sample, and  
6           a fiber optic multiplexer which couples the detection and analysis mechanism to a  
7           second end of each of the plurality of fiber optic cables.

1       16. 4     The apparatus according to claim 13 wherein the sample holder includes a  
2           removable reaction chamber for holding the sample.

1       17.     The apparatus according to claim 16 wherein the removable reaction chamber is  
2           sealable.

1       18. 6     The apparatus according to claim 13 wherein the sample holder includes a sealable  
2           reaction chamber for holding the sample.

1       19. 7     The apparatus according to claim 13 wherein the sample holder includes an optical  
2           interface through which the excitation beam is transmitted from the lens into the sample.

1       20. 8     The apparatus according to claim 19 wherein the sample holder includes a sealable  
2           reaction chamber for holding the sample, the optical interface forming a wall of the reaction  
3           chamber.

1       21. 9     The apparatus according to claim 19 wherein the apparatus further includes a  
2           mechanism for heating the optical interface to prevent condensation of the sample on the  
3           optical interface.

1       22. 10     The apparatus according to claim 21 wherein the sample holder includes a sealable  
2           reaction chamber for holding the sample, the optical interface forming a wall of the reaction  
3           chamber.

1       23. 11     The apparatus according to claim 19 wherein the sample holder includes a

2 removable reaction chamber for holding the sample, the optical interface forming a wall of  
3 the reaction chamber which covers at least a portion of the sample and which is separated  
4 from the sample by an air gap.

1 24. A method for monitoring the formation of a nucleic acid amplification reaction product  
2 in real time comprising:  
3 adding a sample to a sample holder which contains a nucleic acid sequence to be  
4 amplified,  
5 transmitting an excitation beam into the sample which illuminates a volume of the  
6 sample, the sample including a first fluorescent indicator which produces a first fluorescent  
7 signal when illuminated by the excitation beam whose intensity is proportional to the  
8 concentration of amplification reaction product in the sample and the volume of the sample  
9 illuminated by the excitation beam, and a second fluorescent indicator homogeneously  
10 distributed throughout the sample which produces a second fluorescent signal when  
11 illuminated by the excitation beam whose intensity is proportional to the volume of the  
12 sample illuminated by the excitation beam; and  
13 measuring the intensities of the first and second fluorescent signals.

1 25. The method according to claim 24 wherein the first and second fluorescent signals  
2 each have an intensity and the detection, the step of measuring the intensities of the first  
3 and second fluorescent signals including calculating a ratio between the intensity of the first  
4 fluorescent signal and the intensity of the second fluorescent signal.

1 26. The method according to claim 24 wherein the first fluorescent indicator is a  
2 complex-forming dye.

1 27. The method according to claim 24, further including the step of sealing the sample  
2 within the sample holder prior to transmitting an excitation beam into the sample.

1 28. The method according to claim 24 wherein the sample holder includes an optical  
2 interface through which the excitation beam is transmitted from the lens to the sample,

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3 sample holder also including an air gap separating the optical interface from the sample, the  
4 method further including the step of heating the optical interface to prevent condensation of  
5 the sample on the optical interface.

16 15  
1 29. The method according to claim 28, further including the step of sealing the sample  
2 within the sample holder prior to transmitting an excitation beam into the sample.

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1 30. The method according to claim 24 wherein the step of adding a sample to a sample  
2 holder includes  
3 adding a sample to a reaction chamber which is removable from the sample holder;  
4 and  
5 adding the removable reaction chamber to the sample holder.

18 17  
1 31. The method according to claim 30, further including the step of sealing the sample  
2 within the removable reaction chamber.

1 32. 19 17  
The method according to claim 30 wherein the removable reaction chamber includes  
2 an optical interface through which the excitation beam is transmitted from the lens to the  
3 sample and an air gap separating the optical interface from the sample, the method further  
4 including the step of heating the optical interface to prevent condensation of the sample on  
5 the optical interface.

1 33. The method according to claim 24 wherein the nucleic acid amplification reaction is a  
2 polymerase chain reaction.

1 34. The method according to claim 24 wherein the nucleic acid amplification reaction is a  
2 ligase chain reaction.

1 35. The method according to claim 24 wherein the nucleic acid amplification reaction is a  
2 polymerase chain reaction and wherein the first and second fluorescent indicators are  
3 covalently attached to an oligonucleotide having a nucleotide sequence complementary to a

4 portion of a strand of the amplification reaction product, the second fluorescent indicator  
5 quenching the fluorescence of the first fluorescent indicator.

1 36. A method for monitoring the formation of nucleic acid amplification reaction products  
2 in a plurality of samples in real time comprising:  
3 adding samples containing a nucleic acid sequence to be amplified to a plurality of  
4 sample holders;  
5 transmitting excitation beams into the plurality of sample holders which illuminate a  
6 volume of each sample, each sample including a first fluorescent indicator which produces a  
7 first fluorescent signal when illuminated by the excitation beam whose intensity is  
8 proportional to the concentration of amplification reaction product in the sample and the  
9 volume of the sample illuminated by the excitation beam, and a second fluorescent indicator  
10 homogeneously distributed throughout the sample which produces a second fluorescent  
11 signal when illuminated by the excitation beam whose intensity is proportional to the volume  
12 of the sample illuminated by the excitation beam; and  
13 measuring the intensities of the first and second fluorescent signals of each of the  
14 samples.

1 37. The method according to claim 36 wherein at least two different first fluorescent  
2 indicators having different first fluorescent signals are used amongst the plurality of  
3 samples, the step of measuring the intensity of the first fluorescent signal including  
4 measuring the different first fluorescent signals of the at least two different first fluorescent  
5 indicators.

1 38. A method for monitoring the formation of a plurality of nucleic acid amplification  
2 reaction products in a sample in real time comprising:  
3 adding to a sample holder a sample containing a plurality of different nucleic acid  
4 sequences to be amplified,  
5 transmitting an excitation beam into the sample which illuminates a volume of the  
6 sample, the sample including a plurality of first fluorescent indicators which each produce a

7 first fluorescent signal when illuminated by the excitation beam whose intensity is  
8 proportional to the concentration of a particular amplification reaction product in the sample  
9 and the volume of the sample illuminated by the excitation beam, and a second fluorescent  
10 indicator homogeneously distributed throughout the sample which produces a second  
11 fluorescent signal when illuminated by the excitation beam whose intensity is proportional to  
12 the volume of the sample illuminated by the excitation beam; and  
13 measuring the intensities of the first and second fluorescent signals. --.

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